



Process Approach to Detecting Early Cognitive Impairment

Rhoda Au, Ph.D., MBA Friday Harbor 9/14/15

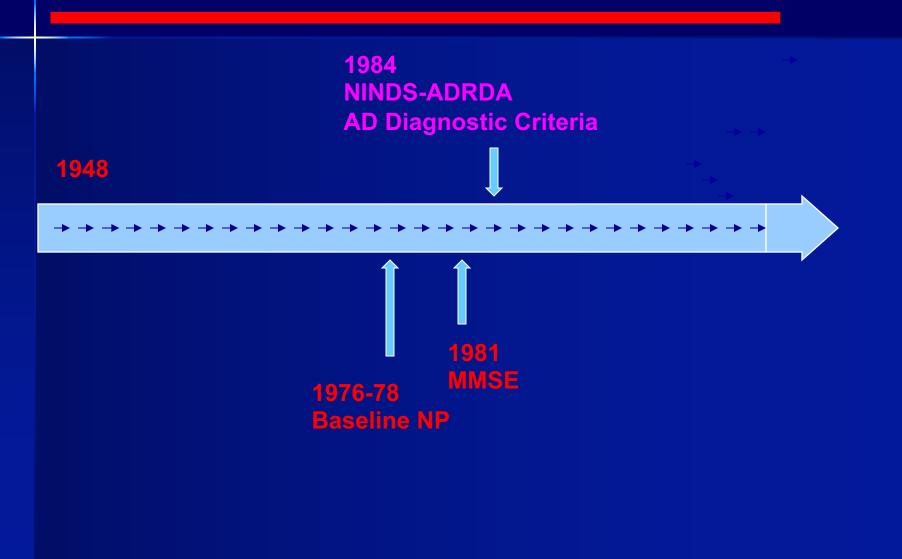
Objectives

Understanding of full scope of FHS cognitive data

 How neuropsychological tests are a richer resource than they are typically used

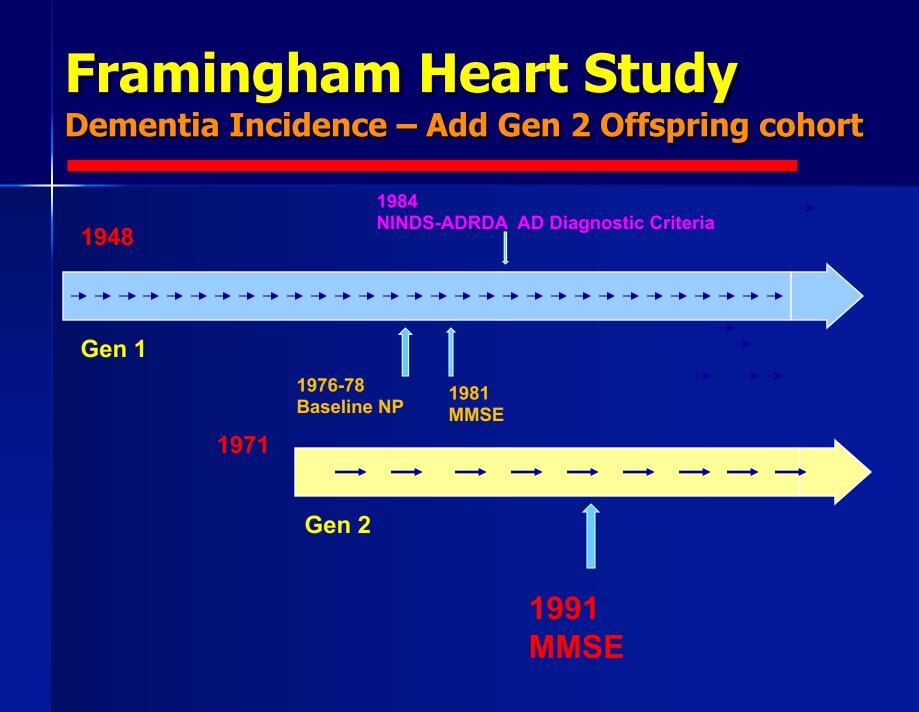
Preview of things to come

Framingham Heart Study Dementia Study – Prevalence: Gen 1 Original cohort

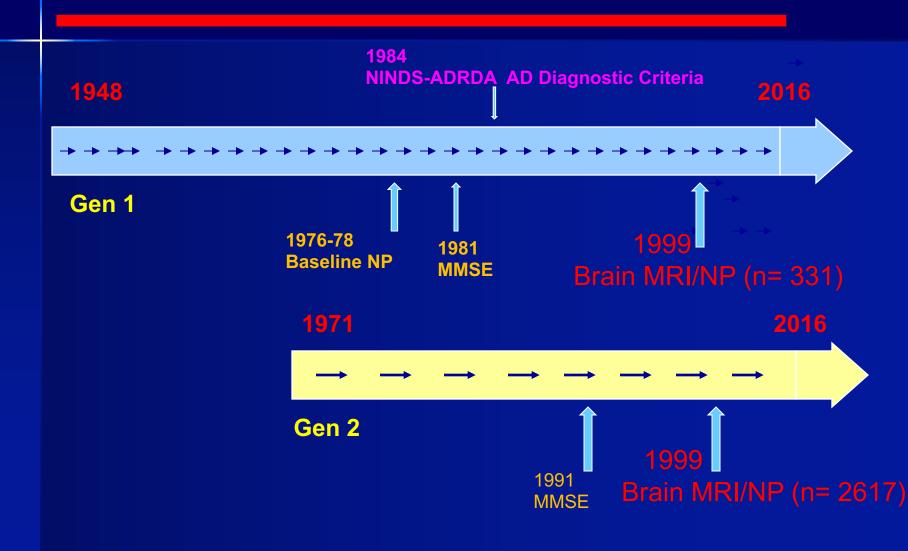


Cogniitve Measures – 1976-78 Gen 1 - Exam 14/15

WMS Logical Memory – IR & DR WMS Visual Reproductions - IR WMS Paired Associates - IR WMS Digit Span WAIS Similarities Controlled Word Association Test - FAS



Framingham Heart Study Incident Dementia + Pre-Clinical



Cognitive Tests - 1999-2005 Gen 1 – Exams 25-27 Gen 2 – Exam 7

WMS Logical Memory – IR & DR WMS Visual Reproductions – IR & DR WMS Paired Associates – IR & DR WAIS Similarities **Digit Span** Controlled Word Association Test -Trails A & B Boston Naming Test – 30 Item **Hooper Visual Organization Test** Finger Tapping WRAT III Reading Subtest

Mild Cognitive Impairment

1999: Preclinical AD

2004: Amnestic Memory Only – Verbal and/or Visual Memory + Other Cognitive Domains

Non-Amnestic Single Domain Multiple Domains

Cognitive Impairment Detection Challenge: FHS Current Two-Method Approach

The Traditional Approach

Boston Process Approach
 Preserve traditional measures

Expand test beyond single score

Focus on path to final response
 Value of incorrect responses

Cognitive Tests – 2005-2011 Gen 1 – Exam 28 Gen 2 – Exam 8

WMS Logical Memory – IR & DR & Recogn WMS Visual Reproductions – IR & DR & Recogn WMS Paired Associates – IR & DR & Recogn WAIS Similarities **Digit Span** Controlled Word Association Test – FAS **Category Fluency - Animals** Trails A & B Boston Naming Test – 30 Item **Hooper Visual Organization Test Finger Tapping** WRAT III Reading Subtest **Clock Drawing Test** WISC-III Math Fluency **Digit Symbol Substitution Test**

Quantitative vs. Qualitative: LM –IR

Traditional scores

Total verbatim Total paraphrase Total – V+P

Qualitative Scores

Confabulations related Intrusions related

Confabulations unrelated Intrusions unrelated

Total Number of Qualitative Measures = 10

Quantitative vs. Qualitative: LM – DR

Qualitative measures from IR PLUS:

Confabulations Related & Unrelated from IR

Related & Unrelated - New

Intrusions Related & Unrelated from IR Related & Unrelated - New

Total Number of Qualitative Measures = 14

Digit Span – Qualitative Errors

Differentiate between sequencing and non-sequencing errors

Sequencing Error example:

For 1 - 5 - 2 - 8 - 6, participant responds 6 - 8 - 2 - 1 - 5

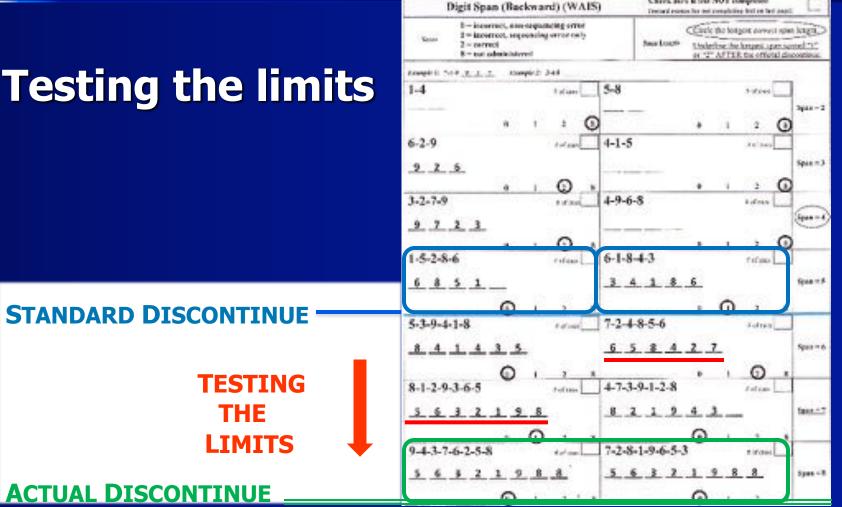
Non-sequencing Error examples:

For 1 - 5 - 2 - 8 - 6, participant responds 6 - 8 - 2 - 5

For 1 - 5 - 2 - 8 - 6, participant responds 6 - 3 - 2 - 1 - 5

Digit Span – Limit Testing

Testing the limits



Chevic here if test NOT completed

Digit Span Calculating Qualitative Scores

Calculate total score accounting for continuum of error types

> Score each trial [0, 1, or 2]
> Assign qualitative value for each digit span [scale 0 to 5]
> Sum qualitative values for qualitative total score

Digit Span – Scoring Trials

TRIAL SCORING:

- -2 = correct response
- -1 = incorrect response
 (sequencing error)
- -0 = incorrect response (nonsequencing error)
- -8 = not administered

Digit Span (Backward) (W.	Clerk kerr if he MDT coupled port case to an majoring out a her say Space reps. Under the server, server can be say Under the server, server can be say Under the driving of part says 1.11 with the server server can be say				
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Digit Span – Scoring Spans

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1	0	1	8-1-2-9-3-6-5 1 due 4-7-3-9-1-2-8 rdue	
0	1	1		1
0	0	0	9.4.3.7.6.2.5.8 5.6.3.2.1.9.8.8 (0 1 3 5 6 3 2 1 9 8 8 (0 1 3 5 6 3 2 1 9 8 8 (0 1 3 5 6 3 2 1 9 8 8 (1 2 8 8 8 1 9 8 8 1 9 8 8 1 9 1 9 1 9 1 9 1	Q

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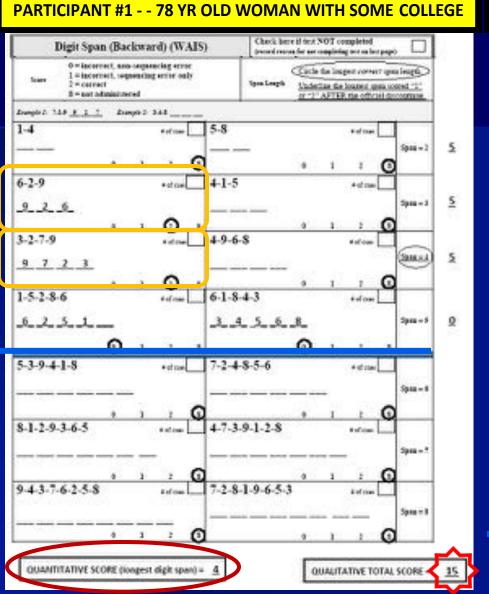
Digit Span (Backward) (WAIS)

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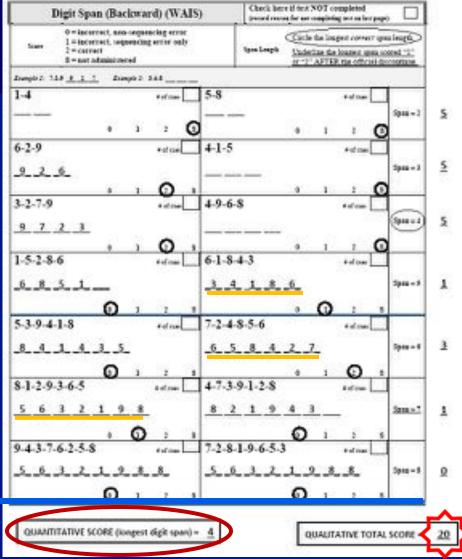
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PARTICIPANT #2 - - 78 YR OLD WOMAN WITH SOME COLLEGE



Qualitative errors – Trails B

Perceptual Errors

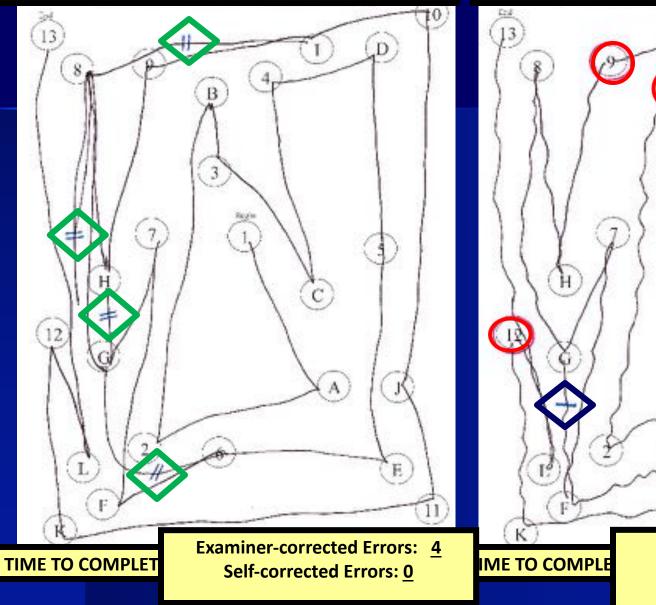
Other Errors

Self-corrected & Examiner-corrected

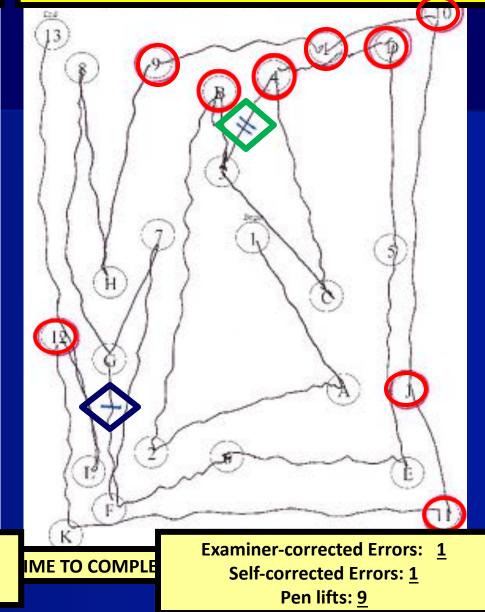
Pen lifts

Starts before told to begin

PARTICIPANT #1 - - 77 YR OLD WOMAN HIGH SCHOOL GRADUATE



PARTICIPANT #2 - - 80 YR OLD WOMAN HIGH SCHOOL GRADUATE



Qualitative errors Verbal fluency



Wrong first letter

Broken rules

Perseverations

ANIMALS

Broken rules

Perseverations

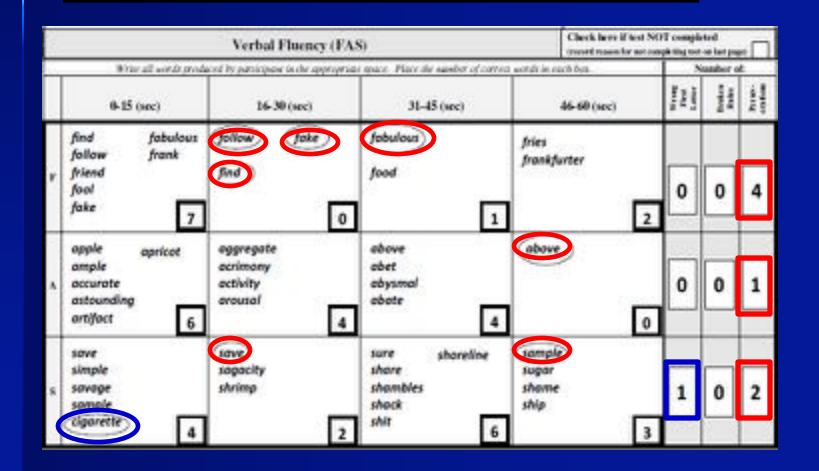
Verbal Fluency

PARTICIPANT #1 - 78 YR OLD MAN Total FAS Score = 39

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Verbal Fluency

PARTICIPANT #2 - 81 YR OLD MAN Total FAS Score = 39



Category Fluency - Animals

64-year old woman

- Quantitative- 15 responses
- Qualitative- 0 errors

ANMALS	cat dog elephant tiger puma horse 7	pony hippopotamus fly 3	mosquito snake 2	bird cow pig 3	X X X	0	0
NN	puma	3	2	3	X	<u> </u>	

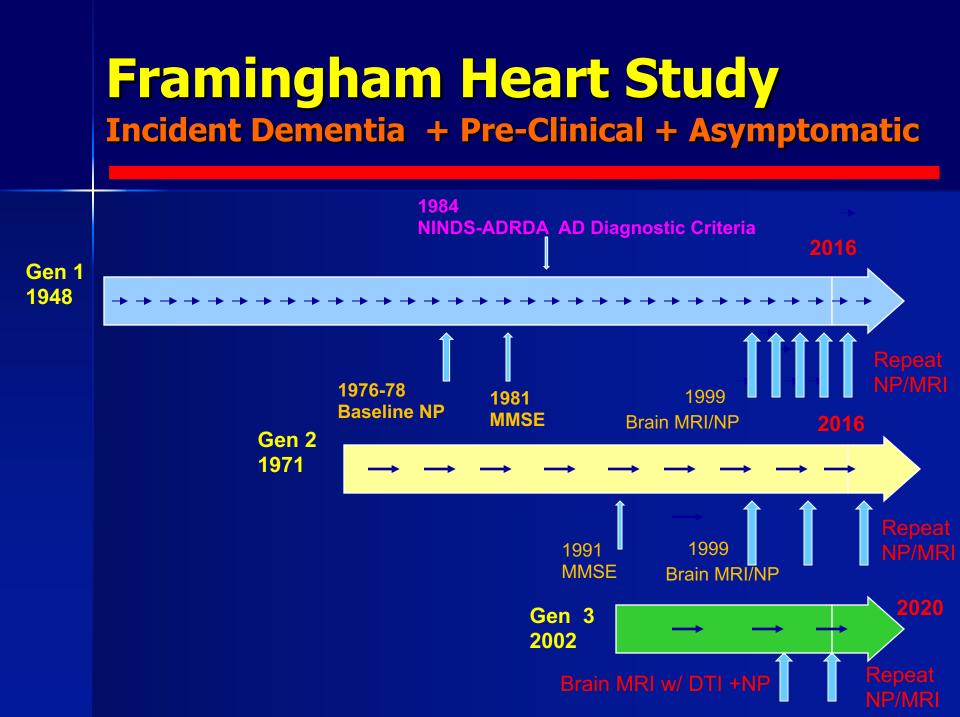
62-year old man

- Quantitative- 15 responses
- Qualitative- 3 errors

ANMALS	cat dog elephant osprey porcupine skunk	antelope deer alligator crocodile koala bear	bear beaver cockatoo	alligator	X X X 1 X 0 3
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Process Across Tests: Executive Function - Perseverations Across Tests

Logical memory - # perseverations: IR & DR Visual Reproductions – perseverations (Y/N per design: IR & DR Paired Associates – perseveration/pair: IR Similarities – perseveration/item FAS - # of perseverations/trial Animals - # of perseverations BNT – perseveration/item



What is Pre-clinical?

65+ years old

Measures differentiate cognitively intact vs. clinically demented

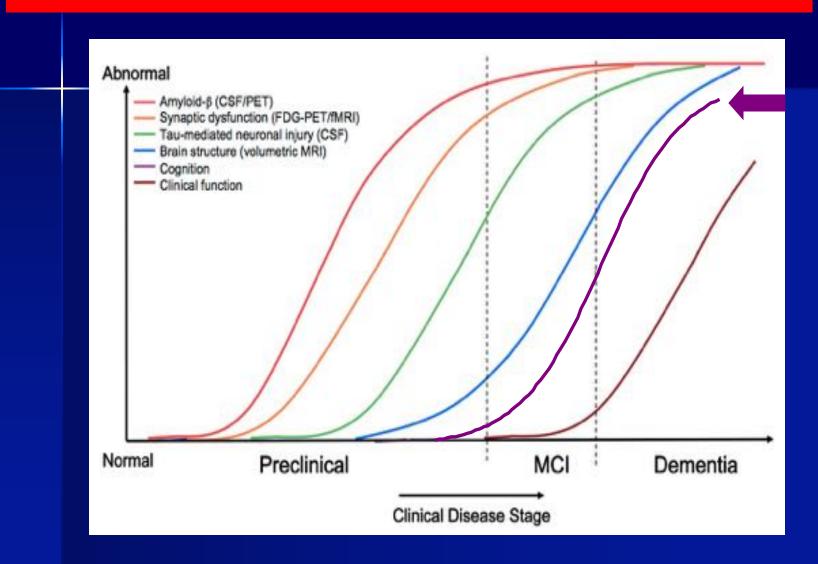
45-65 years old

Measures differentiate cognitively intact vs. pre-clinical

<45 years old</p>

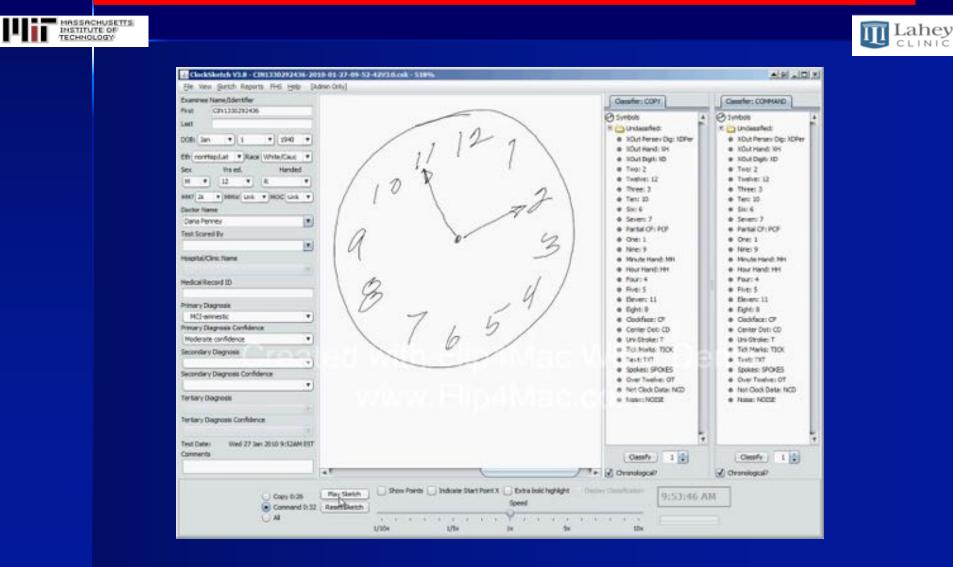
 Measures differentiate cognitively intact vs. cognitively intact

Cognitive Impairment Detection Challenge: When Can It Be Detected?



Adapted by Drs. Dana Penney and Randall Davis from Fig 3 of *Criteria For Preclinical Alzheimer's Disease*, Alzheimer's Association report (2010), which in turn cites Jack C R, et al., Hypothetical model of dynamic biomarkers of the Alzheimer's pathological cascade, *The Lancet*, **9**:1, Jan 2010, pp 119-128.

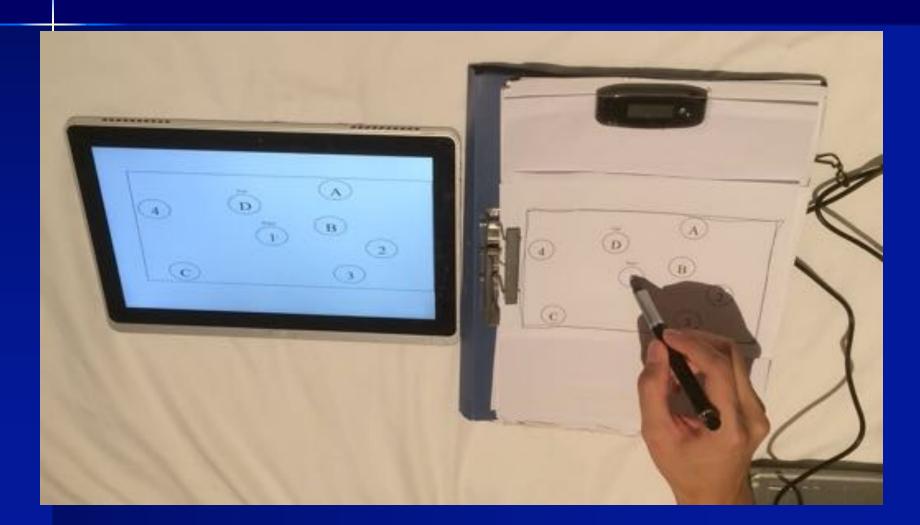
BPA on Steroids Digital Clock Drawing Test - 2011





e-NP Platform





FHS Cognitive Data Today

 Initial BPA data collection in Gen 3 (2009-2013)
 Repeat BPA data collection in Gen 2 (2010-2016)

- Repeat BPA in Gen 3 (2015 2020)
- Digital Ink technology (2011-2020)
- e-NP Platform (2016 2020)